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TITLE: Stress protein-peptide complexes as prophylactic and therapeutic vaccines against intracellular pathogens

CLAIMS:

- 1. A vaccine for administration to a mammal for inducing in the mammal a cytotoxic T cell response against a preselected intracellular pathogen, the vaccine comprising: (a) an immunogenic stress protein-peptide <u>complex</u> operative to initiate in said mammal a cytotoxic T cell response against said pathogen, said <u>complex</u> comprising, a peptide that is present in a eukaryotic cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen, <u>complexed</u> with a stress protein; and (b) a pharmaceutically acceptable carrier.
- 2. A vaccine for administration to a mammal for inducing in said mammal resistance to infection by a preselected intracellular pathogen, the vaccine comprising: (a) an immunogenic stress protein-peptide <u>complex</u> operative to initiate in said mammal, by means of a cytotoxic T cell response in said mammal, resistance to infection by said pathogen, said <u>complex</u> comprising, a peptide that is present in a eukaryotic cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen, <u>complexed</u> with a stress protein; and (b) a pharmaceutically acceptable carrier.
- 3. The composition of claim 1 or 2, wherein said stress protein is a member of the stress protein families selected from the group consisting of Hsp60, Hsp70, and Hsp90.
- 17. The composition of claim 1 or 2, wherein said peptide is non covalently <u>complexed</u> with said stress protein.
- 18. A method of inducing in a mammal a cytotoxic T cell response against a preselected intracellular pathogen that causes disease in said mammal, the method comprising: administering to said mammal a vaccine comprising, (a) an immunogenic stress protein-peptide <u>complex</u> operative to initiate in said mammal a cytotoxic T cell response against said pathogen and comprising, a peptide that is present in a eukaryotic cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen, <u>complexed</u> with a stress protein, and (b) a pharmaceutically acceptable carrier, in an amount sufficient to elicit in said mammal a cytotoxic T cell response against said pathogen.
- 19. A method of inducing in a mammal resistance to infection by a preselected intracellular pathogen that-causes disease in said mammal, the method comprising: administering to said mammal a vaccine comprising, (a) an immunogenic stress protein-peptide complex operative to initiate in said mammal cytotoxic T cell response against said pathogen and comprising, a peptide that is present in a eukaryotic cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen, complexed with a stress protein, and (b) a pharmaceutically acceptable carrier, in an amount sufficient to induce in said mammal, by means of the cytotoxic T cell response in said mammal, resistance to infection by said pathogen.
- 20. The method of claim 18 or 19, wherein said cytotoxic T cell response is mediated by the class I major histocompatibility complex.
- 21. The method of claim 18 or 19, wherein said stress protein is a member of the stress protein families selected from the group consisting of <u>Hsp60</u>, <u>Hsp70</u>, and <u>Hsp90</u>.
- 29. The method of claim 18 or 19, wherein said vaccine is administered to said mammal in an amount in

the range of about 0.1 to about 1000 micrograms of complex/kg body weight of mammal/immunization.

- 30. The method of claim 29, wherein said amount is in range of about 0.5 to about 100 micrograms of <u>complex/kg</u> body weight of mammal/immunization.
- 31. A method for preparing a vaccine for inducing in a mammal a cytotoxic T cell response against a preselected intracellular pathogen, the method comprising: (a) harvesting from a eukaryotic cell infected with said pathogen an immunogenic stress protein-peptide complex comprising, a peptide that is present in said cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen complexed with a stress protein, said complex, when administered to said mammal, being operative at initiating in said mammal a cytotoxic T cell response against said pathogen; and (b) combining said complex with a pharmaceutically acceptable carrier.
- 32. A method for preparing a vaccine for inducing in a mammal a cytotoxic T cell response against a preselected intracellular pathogen, the method comprising: (a) reconstituting in vitro, a peptide that is present in a eukaryotic cell infected with said pathogen but not present in said cell when said cell is not infected with said pathogen and a stress protein, thereby to generate a stress protein-peptide <u>complex</u>, which when administered to said mammal is operative to initiate a cytotoxic T cell response against said pathogen in said mammal; and (b) combining said <u>complex</u> with a pharmaceutically acceptable carrier.
- 33. The method of claim 32, wherein said stress protein is harvested in the presence of <u>ATP</u> prior to reconstitution.
- 35. The method of claim 31 or 32, wherein said stress protein is a member of the stress protein families selected from the group consisting of <u>Hsp60</u>, <u>Hsp70</u> and <u>Hsp90</u>.